



FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 0609.4730000/LBB/M-G	APPLICATION NO. 09/869,565
	APPLICANT Gardella et al.	
	FILING DATE October 17, 2001	GROUP 1643 1646

U.S. PATENT DOCUMENTS							
EXAMINER INITIAL		DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
ZM ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓ ↓	AA1	4,086,196	04/25/1978	Tregear			03/28/1975
	AB1	5,217,896	06/08/1993	Kramer et al.			12/30/1988
	AC1	6,417,333 B1	07/09/2002	Bringhurst et al.			11/24/1999
	AD1	6,495,662 B1	12/17/2002	Gardella et al.			10/20/1999
	AE1	6,537,965 B1	03/25/2003	Bringhurst et al.			11/23/1999
	AF1	6,541,220 B1	04/01/2003	Juppner et al.			11/30/1999
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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
ZM ZM ZM ZM ZM ZM	AL1	WO 87/01130	02/26/1987	WIPO			Yes No
	AM1	WO 91/05050	04/18/1991	WIPO			Yes No
	AN1	EP 0 464 533	01/08/1992	Europe			Yes No
	AO1	EP 0 477 885	04/01/1992	Europe			Yes No
	AP1	WO 92/17602	10/15/1992	WIPO			Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

ZM ↓ ↓ ↓	AR	1	Abou-Samra, A.B., et al., "Phorbol 12-Myristate 13-Acetate and Vasopressin Potentiate the Effect of Corticotropin-releasing Factor on Cyclic AMP Production in Rat Anterior Pituitary Cells," J. Biol. Chem. 262:1129-1136, The American Society for Biochemistry and Molecular Biology, Inc. (1987)
	AS	1	Abou-Samra, A.B., et al., "Cyclic Adenosine 3',5'-Monophosphate (cAMP)-Dependent and cAMP-Independent Regulation of Parathyroid Hormone Receptors on UMR 106-01 Osteoblastic Osteosarcoma Cells," Endocrinology 129:2547-2554, The Endocrine Society (1991)
	AT	1	Abou-Samra, A.B., et al., "Non-Homologous Sequences of Parathyroid and the Parathyroid Hormone Related Peptide Bind to a Common Receptor on ROS 17/2.8 Cells," Endocrinology 125:2215-2217, The Endocrine Society (1989)

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ZH	AL2	EP 0 561 412	09/22/1993	Europe			Yes No
ZH	AM2	GB 2 269 176	02/02/1994	Great Britain			Yes No
ZH	AN2	WO 95/11988	05/04/1995	WIPO			Yes No
ZH	AO2	CA 2,126,132	12/18/1995	Canada			Yes No
ZH	AP2	CA 2,126,299	12/21/1995	Canada			Yes No

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ZH	AR	2	Azarani, A., et al., "Structurally Diverse N-terminal Peptides of Parathyroid Hormone (PTH) and PTH-related Peptide (PTHrP) Inhibit the Na ⁺ /H ⁺ Exchanger NHE3 Isoform by Binding to the PTH/PTHrP Receptor Type I and Activating Distinct Signaling Pathways," J. Biol. Chem. 271:14931-14936, The American Society for Biochemistry and Molecular Biology, Inc. (1996)
	AS	2	Barbier, J.-R., et al., "Bioactivities and Secondary Structures of Constrained Analogues of Human Parathyroid Hormone: Cyclic Lactams of the Receptor Binding Region," J. Med. Chem. 40:1373-1380, American Chemical Society (April 1997)
ZH	AT	2	Bennett, D., et al., "Kinetic Characterization of the Interaction of Biotinylated Human Interleukin 5 with an Fc Chimera of its Receptor α Subunit and Development of a ELISA Screening Assay using Real-Time Interaction Biosensor Analysis," J. Molec. Recognition 8:52-58, John Wiley & Sons, Ltd. (1995)

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ZM ↓ ZM	AL3	EP 0 748 817	12/18/1996	Europe			Yes No
	AM3	WO 97/02834	01/30/1997	WIPO			Yes No
	AN3	WO 98/05683	02/12/1998	WIPO			Yes No
	AO3	WO 99/18945	04/22/1999	WIPO			Yes No
	AP3	WO 00/31137	06/02/2000	WIPO			Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

ZM ↓ ✓	AR	3	Bergwitz, C., et al., "Full Activation of Chimeric Receptors by Hybrids between Parathyroid Hormone and Calcitonin," J. Biol. Chem. 271:26469-26472, The American Society for Biochemistry and Molecular Biology, Inc. (1996)
	AS	3	Bergwitz, C., et al., "Residues in the Membrane-spanning and Extracellular Loop Regions of the Parathyroid Hormone (PTH)-2 Receptor Determine Signaling Selectivity for PTH and PTH-related Peptide," J. Biol. Chem. 272:28861-28868, The American Society for Biochemistry and Molecular Biology, Inc. (November 1997)
	AT	3	Bisello, A., et al., "Parathyroid Hormone-Receptor Interactions Identified Directly by Photocross-linking and Molecular Modeling Studies," J. Biol. Chem. 273:22498-22505, The American Society for Biochemistry and Molecular Biology, Inc. (August 1998)

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EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
3H	AL4	WO 00/32775	06/08/2000	WIPO			Yes No
↓	AM4	WO 00/39278	07/06/2000	WIPO			Yes No
↓	AN4	WO 01/23427	04/05/2001	WIPO			Yes No
↓	AO4	WO 01/23521	04/05/2001	WIPO			Yes No
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)			
3H	AR	4	Bowie, J.U., et al., "Deciphering the Message in Protein Sequences: Tolerance to Amino Acid Substitutions," <i>Science</i> 247:1306-1310, American Association for the Advancement of Science (1990)
↓	AS	4	Caulfield, M.P., et al., "The Bovine Renal Parathyroid Hormone (PTH) Receptor Has Equal Affinity for Two Different Amino Acid Sequences: The Receptor Binding Domains of PTH and PTH-Related Protein Are Located within the 14-34 Region" <i>Endocrinology</i> 127:83-87, The Endocrine Society (1990)
	AT	4	Chorev, M., et al., "Modifications of Position 12 in Parathyroid Hormone and Parathyroid Hormone Related Protein: Toward the Design of Highly Potent Antagonists," <i>Biochem.</i> 29:1580-1586, American Chemical Society (1990)

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	AL						Yes No
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3M	AR	5	Cleland, J.L., et al., "The Development of Stable Protein Formulations: A Close Look at Protein Aggregation, Deamidation, and Oxidation," Crit. Rev. Ther. Drug Carrier Systems 10:307-377, CRC Press, Inc. (1993)
↓	AS	5	Cohen, P.E., et al., "Analogues of Parathyroid Hormone Modified at Positions 3 and 6. Effects on Receptor Binding and Activation of Adenylyl Cyclase in Kidney and Bone," J. Biol. Chem. 266:1997-2004, The American Society for Biochemistry and Molecular Biology, Inc. (1991)
✓	AT	5	Cunningham, B.C., and Wells, J.A., "High-Resolution Epitope Mapping of hGH-Receptor Interactions by Alanine-Scanning Mutagenesis," Science 244:1081-1085, American Association for the Advancement of Science (1989)

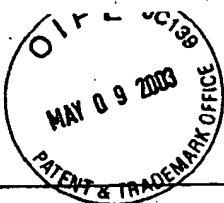
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	AR	6	Dautzenberg, F., et al., "Mapping of the ligand-selective domain of the <i>Xenopus laevis</i> corticotropin-releasing factor receptor 1: Implications for the ligand-binding site," <i>Proc. Natl. Acad. Sci. USA</i> 95:4941-4946, National Academy of Sciences (April 1998)
	AS	6	de Vos, A.M., et al., "Human Growth Hormone and Extracellular Domain of Its Receptor: Crystal Structure of the Complex," <i>Science</i> 255:306-312, American Association for the Advancement of Science (1992)
	AT	6	DeAlmeida, V., and Mayo, K., "Identification of Binding Domains of the Growth Hormone-Releasing Hormone Receptor by Analysis of Mutant and Chimeric Receptor Proteins," <i>Mol. Endocrinol.</i> 12:750-765, The Endocrine Society (May 1998)

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AL						Yes No
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34	AR	2	Dempster, D.W., et al., "Anabolic Actions of Parathyroid Hormone on Bone," Endocr. Rev. 14:690-709, The Endocrine Society (1993)
↓	AS	2	Donahue, H.J., et al., "Differential Effects of Parathyroid Hormone and Its Analogues on Cytosolic Calcium Ion and cAMP Levels in Cultured Rat Osteoblast-like Cells," J. Biol. Chem. 263:13522-13527, The American Society for Biochemistry and Molecular Biology, Inc. (1988)
✓	AT	2	Gardella, T.J., et al., "Mutational Analysis of the Receptor-activating Region of Human Parathyroid Hormone," J. Biol. Chem. 266:13141-13146, The American Society for Biochemistry and Molecular Biology, Inc. (1991).

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34	AR	8	Gardella, T.J., et al., "Determinants of [Arg ²]PTH-(1-34) Binding and Signaling in the Transmembrane Region of the Parathyroid Hormone Receptor," <i>Endocrinology</i> 135:1186-1194, The Endocrine Society (1994)
↓	AS	8	Gardella, T.J., et al., "Analysis of Parathyroid Hormone's Principal Receptor-Binding Region by Site-Directed Mutagenesis and Analog Design," <i>Endocrinology</i> 132:2024-2030, The Endocrine Society (1993)
↓	AT	8	Gardella, T.J., et al., "Transmembrane Residues of the Parathyroid Hormone (PTH)/PTH-related Peptide Receptor That Specifically Affect Binding and Signaling by Agonist Ligands," <i>J. Biol. Chem.</i> 271:12820-12825, The American Society for Biochemistry and Molecular Biology, Inc. (1996)

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34	AR	2	Gardella, T.J., et al., "Parathyroid Hormone (PTH)-PTH-related Peptide Hybrid Peptides Reveal Functional Interactions between the 1-14 and 15-34 Domains of the Ligand," J. Biol. Chem. 270:6584-6588, The American Society for Biochemistry and Molecular Biology, Inc. (1995)
↓	AS	2	Gentz, R., et al., "Bioassay for trans-activation using purified human immunodeficiency virus tat-encoded protein: Trans-activation requires mRNA synthesis," Proc. Natl. Acad. Sci. USA 86:821-824, National Academy of Sciences (1989)
↓	AT	2	Gombert, F., et al., "Alanine and D-Amino Acid Scan of Human Parathyroid Hormone," in Peptides: Chemistry, Structure and Biology. Proceedings of the 14th American Peptide Symposium, June 18-23, Kaumaya, P.T.P., and Hodges, R.S., eds., Mayflower Scientific Ltd., Kingswinford, UK, pp. 661-662 (1996)

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34	AR	10	Guo, J., et al., "Parathyroid Hormone (PTH)/PTH-Related Peptide Receptor Density Modulates Activation of Phospholipase C and Phosphate Transport by PTH in LLC-PK1 Cells," <i>Endocrinology</i> 136:3884-3891, The Endocrine Society (1995)
↓	AS	10	Hjorth, S., et al., "Constitutive Activity of Glucagon Receptor Mutants," <i>Mol. Endocrinol.</i> 12:78-86, The Endocrine Society (January 1998)
↓	AT	10	Holtmann, M., et al., "Critical Contributions of Amino-terminal Extracellular Domains in Agonist Binding and Activation of Secretin and Vasoactive Intestinal Polypeptide Receptors. Studies of Chimeric Receptors," <i>J. Biol. Chem.</i> 270:14394-14398, The American Society for Biochemistry and Molecular Biology, Inc. (1995)

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3H	AR	11	Horiuchi, N., et al., "A Parathyroid Hormone Inhibitor in vivo: Design and Biological Evaluation of a Hormone Analog," <i>Science</i> 220:1053-1055, American Association for the Advancement of Science (1983)
↓	AS	11	Hruska, K.A., et al., "Stimulation of Inositol Trisphosphate and Diacylglycerol Production in Renal Tubular Cells by Parathyroid Hormone," <i>J. Clin. Invest.</i> 79:230-239, The Rockefeller University Press (1987)
↓	AT	11	Ji, I.H., and Ji, T.H., "Human Choriogonadotropin Binds to a Lutropin Receptor with Essentially No N-terminal Extension and Stimulates cAMP Synthesis," <i>J. Biol. Chem.</i> 266:13076-13079, The American Society for Biochemistry and Molecular Biology, Inc. (1991)

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34	AR	12	Johanson, K., et al., "Binding Interactions of Human Interleukin 5 with Its Receptor α Subunit. Large Scale Production, Structural, and Functional Studies of <i>Drosophila</i> -Expressed Recombinant Proteins," <i>J. Biol. Chem.</i> 270:9459-9471, The American Society for Biochemistry and Molecular Biology, Inc. (1995)
↓	AS	12	Jüppner, H., et al., "A G Protein-Linked Receptor for Parathyroid Hormone and Parathyroid Hormone-Related Peptide," <i>Science</i> 254:1024-1026, American Association for the Advancement of Science (1991)
↓	AT	12	Jüppner, H., et al., "The Extracellular Amino-Terminal Region of the Parathyroid Hormone (PTH)/PTH-Related Peptide Receptor Determines the Binding Affinity for Carboxyl-Terminal Fragments of PTH-(1-34)," <i>Endocrinology</i> 134:879-884, The Endocrine Society (1994)

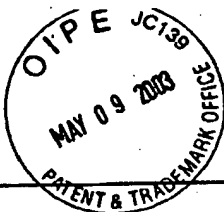
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	AL						Yes No
	AM						Yes No
	AN						Yes No
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34	AR	13	Kolakowski, L.F., "GCRDb: A G-Protein-Coupled Receptor Database," <i>Receptors and Channels</i> 2:1-7, Harwood Academic Publishers (1994)
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3M	AR	15	Mannstadt, M., et al., "Evidence for a Ligand Interaction Site at the Amino-Terminus of the Parathyroid Hormone (PTH)/PTH-related Protein Receptor from Cross-linking and Mutational Studies," J. Biol. Chem. 273:16890-16896, The American Society for Biochemistry and Molecular Biology, Inc. (July 1998)
3M	AS	15	McCuaig, K.A., et al., "Molecular cloning of the gene encoding the mouse parathyroid hormone/parathyroid hormone-related peptide receptor," Proc. Natl. Acad. Sci. USA 91:5051-5055, National Academy of Sciences (1994)
3M	AT	15	McGeoch, D.J., "On the predictive recognition of signal peptide sequences," Virus Res. 3:271-286, Elsevier Science B.V. (1985)

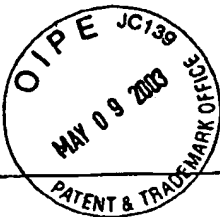
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3H	AT	16	Neugebauer, W., et al., "Solution Structure and Adenylyl Cyclase Stimulating Activities of C-Terminal Truncated Human Parathyroid Hormone Analogues," <i>Biochem.</i> 34:8835-8842, American Chemical Society (1995)

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	AS	17	Nussbaum, S.R., et al., "Parathyroid Hormone•Renal Receptor Interactions. Demonstration of Two Receptor-Binding Domains," J.Biol.Chem. 255:10183-10187. The American Society of Biological Chemists, Inc. (1980)
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3H	AT	18	Rixon, R.H., et al., "Parathyroid Hormone Fragments May Stimulate Bone Growth in Ovariectomized Rats by Activating Adenylyl Cyclase," J. Bone Min. Res. 9:1179-1189, Mary Ann Liebert, Inc. (1994)

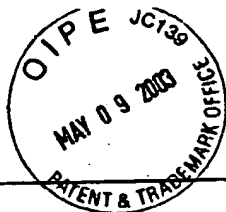
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3H	AR	19	Robbins, D.C., et al., "Antibodies to Covalent Aggregates of Insulin in Blood of Insulin-Using Diabetic Patients," <i>Diabetes</i> 36:838-841, American Diabetes Association (1987)
3H	AS	19	Rubin, D.A., and Jüppner, H., "Zebrafish Express the Common Parathyroid Hormone/Parathyroid Hormone-related Peptide Receptor (PTH1R) and a Novel Receptor (PTH3R) That Is Preferentially Activated by Mammalian and Pufferfish Parathyroid Hormone-related Peptide," <i>J. Biol. Chem.</i> 274:28185-28190, The American Society for Biochemistry and Molecular Biology, Inc. (October 1999)
3M	AT	19	Rosenblatt, M., "Parathyroid Hormone: Chemistry and Structure-Activity Relations," in <i>Pathobiology Annual Vol. 11</i> , Ioachim, H.L., ed., Raven Press, NY, pp. 53-86 (1981)

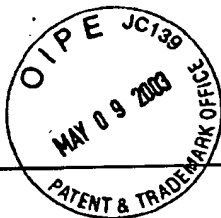
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34	AR	20	Shimizu, M., et al., "Type-Substitution Analysis of the Amino-Terminal Fragment of Parathyroid Hormone, PTH(1-14): An Approach toward New Low Molecular Weight PTH Agonists," J. Bone Min. Res. 14(suppl. 1):Abs. No. F398, Mary Ann Liebert, Inc. (September 1999)
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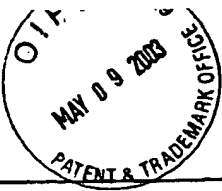
3H	AR	21	Smith, L.J., et al., "Human Interleukin 4. The Solution Structure of a Four-helix Bundle Protein," J. Mol. Biol. 224:899-904, Academic Press Ltd. (1992)
↓	AS	21	Smith, D.B., and Johnson, K.S., "Single-step purification of polypeptides expressed in <i>Escherichia coli</i> as fusions with glutathione S-transferase," Gene 67:31-40, Elsevier Science B.V. (1988)
↓	AT	21	Smith, T.F., and Waterman, M.S., "Comparison of Biosequences," Adv. Applied Math 2:482-489, Academic Press, Inc. (1981)

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↓	AT	22	Szabo, P., "In Situ Hybridization," in <i>Human Chromosomes. Manual of Basic Techniques</i> , Verma, R.S., and Babu, A., eds., Pergamon Press, New York, NY, pp.152-165 (1989)

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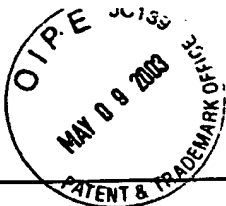
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3H	AT	24	Turner, P.R., et al., "Transmembrane Residues Together with the Amino Terminus Limit the Response of the Parathyroid Hormone (PTH) 2 Receptor to PTH-related Peptide," J. Biol. Chem. 273:3830-3837, The American Society for Biochemistry and Molecular Biology, Inc. (February 1998)

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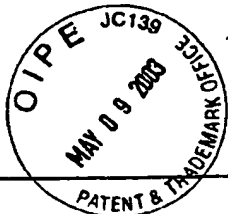
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FORM PTO-1449 INFORMATION DISCLOSURE STATEMENT	ATTY. DOCKET NO. 0609.4730000/LBB/M-G	APPLICATION NO. 09/869,565
	APPLICANT Gardella et al.	
	FILING DATE October 17, 2001	GROUP 1645 1646

U.S. PATENT DOCUMENTS							
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	AL						Yes No
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301	AR	26	Wilson, I.A., et al., "The Structure of an Antigenic Determinant in a Protein," Cell 37:767-778, MIT (1984)
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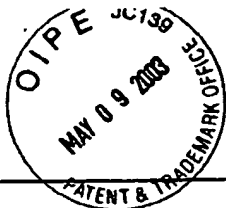
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OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

3M	AR	27	International Search Report for International Patent Application PCT/US99/28207 (mailed April 18, 2000)
3M	AS	27	International Search Report for International Patent Application PCT/US00/04716, (mailed October 11, 2000)
3M	AT	27	Dialog File 351, Accession No. 8882525, Derwent WPI English language abstract for European Patent Publication No. EP 0 464 533 (Document AN1), Derwent Information Ltd. (1992)

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3M	AP4	WO 92/01810	02/06/1992	WIPO			Yes
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AL							Yes No
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3H	AR	29	Broadus, A.E., and Stewart, A.F., "Parathyroid Hormone-Related Protein, Structure, Processing, and Physiological Actions," in <i>The Parathyroids</i> , Bilezikian, J.P., ed., Raven Press, Ltd., New York, NY, pp.259-294 (1994)
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	AN						Yes No
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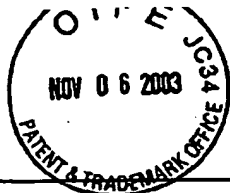
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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

341	AR	33	Orloff, J.J., et al., "A Midregion Parathyroid Hormone-Related Peptide Mobilizes Cytosolic Calcium and Stimulates Formation of Inositol Triphosphate in a Squamous Carcinoma Cell Line," <i>Endocrinol.</i> 137:5376-5385, The Endocrine Society (1996)
↓	AS	33	Orloff, J.J., et al., "Further Evidence for a Novel Receptor for Amino-Terminal Parathyroid Hormone-Related Protein on Keratinocytes and Squamous Carcinoma Cell Lines," <i>Endocrinol.</i> 136:3016-3023, The Endocrine Society (1995)
✓	AT	33	Potts, Jr., J.T., and Juppner, H., "Parathyroid Hormone and Parathyroid Hormone-Related Peptide in Calcium Homeostasis, Bone Metabolism, and Bone Development: The Proteins, Their Genes, and Receptors," in <i>Metabolic Bone Disease</i> , 3 rd Edition, Avioli, L.V., and Krane, S.M., eds., Academic Press, San Diego, CA, pp. 51-94 (1998)

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Gardella et al.FILING DATE
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1645 1646**U.S. PATENT DOCUMENTS**

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	AN						Yes No
	AO						Yes No
	AP						Yes No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

3M	AR	34	Rubin, D.A., et al., "Molecular Cloning and Expression of Receptors for Parathyroid Hormone (PTH) and PTH-Related (PTHrP) Protein in Zebrafish," Am. Zool. 36:97A Abs No. 373, Society for Integrative and Comparative Biology (1996)
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3M	AR	35	Schipani, E., et al., "Pseudohypoparathyroidism Type Ib Is not Caused by Mutations in the Coding Exons of the Human Parathyroid Hormone (PTH)/PTH-Related Peptide Receptor Gene," J. Clin. Endocrinol. Metab. 80:1611-1621, The Endocrine Society (1995)
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U.S. PATENT DOCUMENTS

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3M	AR	36	Takasu, H. et al., "Human PTH/PTHrP receptors and Type-2 PTH Receptors Show Discordant Selectivity for Human PTH Analogs with Amino-Terminal Modifications," <i>Bone</i> 23:S255, Abstract No. T223, Elsevier Science (November 1998)
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	APPLICANT Gardella et al.	
	FILING DATE October 17, 2001	GROUP 1645 1646

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34 ↓ ✓	AR	37	Wu, T.L., et al., "Structural and Physiologic Characterization of the Mid-region Secretory Species of Parathyroid Hormone-related Protein," J. Biol. Chem. 271:24371-24381, The American Society for Biochemistry and Molecular Biology, Inc. (1996)
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FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL		DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
ZH	AL5	WO 00/23594	08/16/2001	WIPO			Yes
	AM5	WO 00/31266	09/19/2001	WIPO			No
	AN5	WO 00/40698	10/10/2001	WIPO			Yes
	AO5	WO 03/009804	02/06/2003	WIPO			No
	AP5	WO 2004/067021	08/12/2004	WIPO			Yes
							No

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

AR	38	U.S. Patent Application No. 09/475,158, Gardella <i>et al.</i> , filed December 30, 1999 (NOT PUBLISHED)
AS	38	U.S. Patent Application No. 09/672,020, Gardella <i>et al.</i> , filed September 29, 2000 (NOT PUBLISHED)
AT	38	U.S. Patent Application No. 10/959,605, Bringhurst <i>et al.</i> , filed October 7, 2004 (NOT PUBLISHED)

EXAMINER

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405802_1.DOC

Considered, do
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FORM PTO-1449

**SECOND SUPPLEMENTAL
INFORMATION DISCLOSURE STATEMENT**

 ATTY. DOCKET NO.
0609.4730000/SRL/PAC

 APPLICATION NO.
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 APPLICANTS
Gardella *et al.*

 FILING DATE
October 17, 2001

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U.S. PATENT DOCUMENTS

EXAMINER INITIAL	DOCUMENT NUMBER	DATE	NAME	CLASS	SUB-CLASS	FILING DATE
AA						
AB						
AC						
AD						
AE						
AF						
AG						
AH						
AI						
AJ						
AK						

FOREIGN PATENT DOCUMENTS

EXAMINER INITIAL	DOCUMENT NUMBER	DATE	COUNTRY	CLASS	SUB-CLASS	TRANSLATION
3M	AL6	WO 2004/093902	11/04/2004	WIPO		Yes
3M	AM6	WO 2005/009358	02/03/2005	WIPO		No
	AN					Yes
	AO					No
	AP					Yes

OTHER (Including Author, Title, Date, Pertinent Pages, etc.)

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AS			
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